



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of:

FIKES *et al.*

Appl. No. 09/458,297

Filed: December 10, 1999

For: **Inducing Cellular Immune
Responses To p53 Using Peptide
and Nucleic Acid Compositions**

Confirmation No. 8696

Art Unit: 1644

Examiner: Schwadron, R.B.

Atty. Docket:
2060.0120000/EKS/HCC/M-M

Second Supplemental Information Disclosure Statement

Commissioner for Patents
PO Box 1450
Alexandria, VA 22313-1450

Sir:

Listed on accompanying Form PTO-1449 are documents that may be considered material to the examination of this application, in compliance with the duty of disclosure requirements of 37 C.F.R. §§ 1.56, 1.97 and 1.98. A copy of each document is also provided.

In accordance with 37 C.F.R. § 1.98(a)(3), Applicants' undersigned representative submits the following, in regards to non-English language documents AM1, AP1, and AM2 cited on Form PTO 1449:

Document AM1, EP 0 226 513, is in the French language. It is believed that AM1 discloses HIV peptide derivatives. An English language abstract of document AM1 is attached as Document AS25;

Document AP1, WO 92/21033, is in the German language. It is believed that AP1 discloses peptides which bind MHC class I and class II molecules. An English language

abstract of document AP1 is located on page one of the document and another English language abstract is attached as Document AT25;

Document AM2, WO 94/11738, is in the German language. It is believed that AM2 discloses peptides which bind MHC class I and class II molecules. An English language abstract of document AM2 is located on page one of the document and another English language abstract is attached as Document AR26.

Where the publication date of a listed document does not provide a month of publication, the year of publication of the listed document is sufficiently earlier than the effective U.S. filing date and any foreign priority date so that the month of publication is not in issue. Applicants have listed publication dates on the attached PTO-1449 based on information presently available to the undersigned. However, the listed publication dates should not be construed as an admission that the information was actually published on the date indicated.

Applicants reserve the right to establish the patentability of the claimed invention over any of the information provided herewith, and/or to prove that this information may not be prior art, and/or to prove that this information may not be enabling for the teachings purportedly offered.

This statement should not be construed as a representation that a search has been made, or that information more material to the examination of the present patent application does not exist. The Examiner is specifically requested not to rely solely on the material submitted herewith.

This Second Supplemental Information Disclosure Statement is being filed before the mailing date of a first Office Action on the merits. No statement or fee is required.

It is respectfully requested that the Examiner initial and return a copy of the enclosed PTO-1449, and indicate in the official file wrapper of this patent application that the documents have been considered.

The U.S. Patent and Trademark Office is hereby authorized to charge any fee deficiency, or credit any overpayment, to our Deposit Account No. 19-0036.

Respectfully submitted,

STERNE, KESSLER, GOLDSTEIN & FOX P.L.L.C.



Helene C. Carlson
Agent for Applicants
Registration No. 47,473

Date: 8/15/03

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Washington, D.C. 20005-3934
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FORM PTO-1449

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FOREIGN PATENT DOCUMENTS

EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB- CLASS	TRANSLATION
	AL1	EP 0 044 710 A1	01/27/1982	Europe			Yes No
	AM1	EP 0 226 513 A1	06/24/1987	Europe			Yes X No
	AN1	WO 92/02543 A1	02/20/1992	WIPO			Yes No
	AO1	WO 92/12996 A2	08/06/1992	WIPO			Yes No
	AP1	WO 92/21033 A1	11/26/1992	WIPO			Yes X No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>1</u>	Aichele, P., et al., "Antiviral cytotoxic T cell response induced by in vivo priming with a free synthetic peptide," <i>J. Exp. Med.</i> 171:1815-1820, Rockefeller University Press (1990)
	AS	<u>1</u>	Alexander, J., et al., "Derivation of HLA-A11/K ^b Transgenic Mice. Functional CTL Repertoire and Recognition of Human A11-Restricted CTL Epitopes," <i>J. Immunol.</i> 159:4753-4761, The American Association of Immunologists (1997)
	AT	<u>1</u>	Bergmann, C.C., et al., "Differential Effects of Flanking Residues on Presentation of Epitopes from Chimeric Peptides," <i>J. Virol.</i> 68:5306-5310, American Society for Microbiology (August 1994)

EXAMINER

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	AL2	WO 93/03764 A1	03/04/1993	WIPO			Yes No
	AM2	WO 94/11738 A1	05/26/1994	WIPO			Yes X No
	AN2	WO 94/20127 A1	09/15/1994	WIPO			Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	2	Bertoni, R., et al., "Human Histocompatibility Leukocyte Antigen-binding Supermotifs Predict Broadly Cross-reactive Cytotoxic T Lymphocyte Responses in Patients with Acute Hepatitis," <i>J. Clin. Invest.</i> 100:503-513, The American Society for Clinical Investigation, Inc. (1997)
	AS	2	Bertoni, R., et al., "Human Class I Supertypes and CTL Repertoires Extend to Chimpanzees," <i>J. Immunol.</i> 161:4447-4455, American Association of Immunologists (1998)
	AT	2	Bjorkman, P.J., et al., "Structure of the human class I histocompatibility antigen, HLA-A2," <i>Nature</i> 329:506-512, Macmillan Publishers, Ltd. (1987)

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	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	3	Bjorkman, P.J., et al., "The foreign antigen binding site and T cell recognition regions of class I histocompatibility antigens," Nature 329:512-518, Macmillan Publishers, Ltd. (1987)
	AS	3	Buus, S., et al., "Autologous Peptides Constitutively Occupy the Antigen Binding Site on Ia," Science 242:1045-1047, American Association for the Advancement of Science (1988)
	AT	3	Carreno, B.M., et al., "HLA-B37 and HLA-A2.1 molecules bind largely nonoverlapping sets of peptides," Proc. Natl. Acad. Sci. USA 87:3420-3424, National Academy Press (1990)

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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>4</u>	Corr, M., et al., "Endogenous Peptides of a Soluble Major Histocompatibility Complex Class I Molecule, H-2L ^d : Sequence Motif, Quantitative Binding, and Molecular Modeling of the Complex," <i>J. Exp. Med.</i> 176:1681-1692, Rockefeller University Press (December 1992)
	AS	<u>4</u>	De Bruijn, M.L.H., et al., "Peptide loading of empty major histocompatibility complex molecules on RMA-S cells allows the induction of primary cytotoxic T lymphocyte responses," <i>Eur. J. Immunol.</i> 21:2963-2970, VCH Verlagsgesellschaft mbH (1991)
	AT	<u>4</u>	Del Val, M., et al., "Efficient Processing of an Antigenic Sequence for Presentation by MHC Class I Molecules Depends on Its Neighboring Residues in the Protein," <i>Cell</i> 66:1145-1153, Cell Press (1991)

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	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>5</u>	Deres, K., et al., "In vivo priming of virus-specific cytotoxic T lymphocytes with synthetic lipopeptide vaccine," <i>Nature</i> 342:561-564, Macmillan Publishers, Ltd. (1989)
	AS	<u>5</u>	DiBrino, M., et al., "HLA-A1 and HLA-A3 T Cell Epitopes Derived from Influenza Virus Proteins Predicted from Peptide Binding Motifs," <i>J. Immunol.</i> 151:5930-5935, The Association of Immunologists (December 1993)
	AT	<u>5</u>	DiBrino, M., et al., "Endogenous peptides bound to HLA-A3 possess a specific combination of anchor residues that permit identification of potential antigenic peptides," <i>Proc. Natl. Acad. Sci. USA</i> 90:1508-1512, National Academy Press (February 1993)

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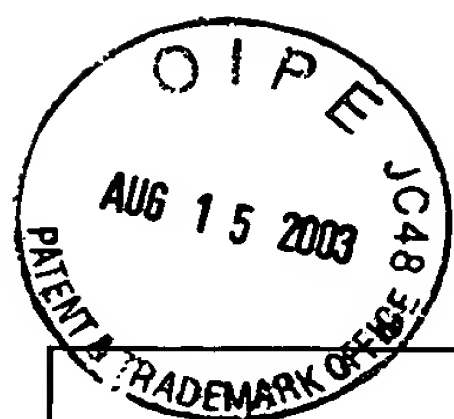
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	AR	<u>6</u>	Ding, Y.-H., et al., "Two Human T Cell Receptors Bind in a Similar Diagonal Mode to the HLA-A2/Tax Peptide Complex Using Different TCR Amino Acids," <i>Immunity</i> 8:403-11, Cell Press (1998)
	AS	<u>6</u>	Eisenlohr, L.C., et al., "Flanking Sequences Influence the Presentation of an Endogenously Synthesized Peptide to Cytotoxic T Lymphocytes," <i>J. Exp. Med.</i> 175:481-487, The Rockefeller University Press (February 1992)
	AT	<u>6</u>	Engelhard, V.H., "Structure of peptides associated with MHC Class I molecules," <i>Curr. Opin. Immunol.</i> 6:13-23, Current Biology, Ltd. (February 1994)

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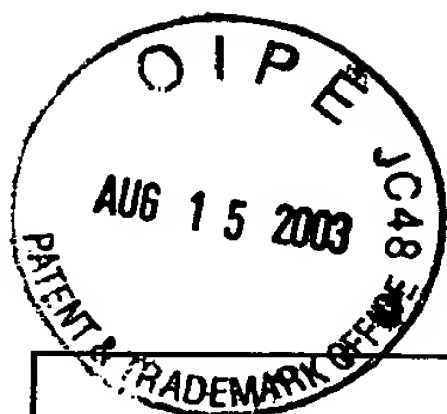
OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>7</u>	Falk, K., et al., "Allele-specific motifs revealed by sequencing of self-peptides eluted from MHC molecules," <i>Nature</i> 351:290-296, Macmillan Publishers, Ltd. (1991)
	AS	<u>7</u>	Falk, K., et al., "MHC peptide motif register. Peptide motifs of HLA-B35 and -B37 molecules," <i>Immunogenetics</i> 38:161-162, Springer-Verlag (April 1993)
	AT	<u>7</u>	Falk, K., et al., "Allele-specific peptide ligand motifs of HLA-C molecules," <i>Proc. Natl. Acad. Sci. USA</i> 90:12005-12009, National Academy Press (December 1993)

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	AR	8	Falk, K., et al., "Pool sequencing of natural HLA-DR, DQ, and DP ligands reveals detailed peptide motifs, constraints of processing, and general rules," <i>Immunogenetics</i> 39:230-242, Springer-Verlag (February 1994)
	AS	8	Falk, K., et al., "Peptide motifs of HLA-A1, -A11, -A31, and -A33 molecules," <i>Immunogenetics</i> 40:238-241, Springer-Verlag (July 1994)
	AT	8	Foon, K.A., "Biological Response Modifiers: The New Immunotherapy," <i>Cancer Res.</i> 49:1621-1639, American Association for Cancer Research (1989)

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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)			
	AR	9	Geysen, H.M., et al., "Cognitive Features of Continuous Antigenic Determinants," J. Mol. Recognit. 1:32-41, Heyden & Sons, Ltd. (1988)
	AS	9	Guo, H.-C., et al., "Different length peptides bind to HLA-Aw68 similarly at their ends but bulge out in the middle," Nature 360:364-366, Macmillan Publishers, Ltd. (November 1992)
	AT	9	Henderson, R.A., et al., "HLA-A2.1-Associated Peptides from a Mutant Cell Line: A Second Pathway of Antigen Presentation," Science 255:1264-1266, American Association for the Advancement of Science (March 1992)

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	AR	<u>10</u>	Hill, A., et al., "Characterization of two Epstein-Barr virus epitopes restricted by HLA-B7," <i>Eur. J. Immunol.</i> 25:18-24, VCH Verlagsgesellschaft mbH (1995)
	AS	<u>10</u>	Hunt, D.F., et al., "Characterization of Peptides Bound to the Class I MHC Molecule HLA-A2.1 by Mass Spectrometry," <i>Science</i> 255:1261-1263, American Association for the Advancement of Science (March 1992)
	AT	<u>10</u>	Ishioka, G.Y., et al., "Utilization of MHC Class I Transgenic Mice for Development of Minigene DNA Vaccines Encoding Multiple HLA-Restricted CTL Epitopes," <i>J. Immunol.</i> 162:3915-3925, The American Association of Immunologists (1999)

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	AR	<u>11</u>	Jameson, S.C., and Bevan, M.J., "Dissection of major histocompatibility complex (MHC) and T cell receptor contact residues in a K ^b -restricted ovalbumin peptide and an assessment of the predictive power of MHC-binding motifs," <i>Eur. J. Immunol.</i> 22:2663-2667, VCH Verlagsgesellschaft mbH (October 1992)
	AS	<u>11</u>	Jardetzky, T.S., et al., "Identification of self peptides bound to purified HLA-B27," <i>Nature</i> 353:326-329, Macmillan Publishers, Ltd. (1991)
	AT	<u>11</u>	Kannagi, M., et al., "Target Epitope in the Tax Protein of Human T-Cell Leukemia Virus Type I Recognized by Class I Major Histocompatibility Complex-Restricted Cells," <i>J. Virol.</i> 66:2928-2933, American Society for Microbiology (May 1992)

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	AR	<u>12</u>	Kast, W.M., et al., "Protection against lethal Sendai virus infection by <i>in vivo</i> priming of virus-specific cytotoxic T lymphocytes with a free synthetic peptide," <i>Proc. Natl. Acad. Sci. USA</i> 88:2283-2287, National Academy Press (1991)
	AS	<u>12</u>	Kast, W.M., et al., "Strict peptide length is not required for the induction of cytotoxic T lymphocyte-mediated antiviral protection by peptide vaccination," <i>Eur. J. Immunol.</i> 23:1189-1192, VCH Verlagsgesellschaft mbH (May 1993)
	AT	<u>12</u>	Krieger, J.I., et al., "Single amino acid changes in DR and antigen define residues critical for peptide-MHC binding and T cell recognition," <i>J. Immunol.</i> 146:2331-2340, American Association of Immunologists (1991)

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	AR	<u>13</u>	Lipford, G.B., et al., "Primary in Vivo Responses to Ovalbumin. Probing the Predictive Value of the K ^b Binding Motif," <i>J. Immunol.</i> 150:1212-1222, The American Association of Immunologists (February 1993)
	AS	<u>13</u>	Maryanski, J.L., et al., "Synthetic peptides as antigens and competitors in recognition by H-2-restricted cytolytic T cells specific for HLA," <i>J. Exp. Med.</i> 167:1391-1405, Rockefeller University Press (1988)
	AT	<u>13</u>	Maryanski, J.L., et al., "Competitor Analogs for Defined T Cell Antigens: Peptides Incorporating a Putative Binding Motif and Polyproline or Polyglycine Spacers," <i>Cell</i> 60:63-72, Cell Press (1990)

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	AL						Yes No
	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>14</u>	Morrison, J., et al., "Identification of the nonamer peptide from influenza A matrix protein and the role of pockets of HLA-A2 in its recognition by cytotoxic T lymphocytes," <i>Eur. J. Immunol.</i> 22:903-907, VCH Verlagsgesellschaft mbH (April 1992)
	AS	<u>14</u>	Niedermann, G., et al., "The proteolytic fragments generated by vertebrate proteosomes: Structural relationships to major histocompatibility complex class I binding peptides," <i>Proc. Natl. Acad. Sci. USA</i> 93:8572-8577, National Academy Press (1996)
	AT	<u>14</u>	Ochoa-Garay, J., et al., "The ability of peptides to induce cytotoxic T cells in vitro does not strongly correlate with their affinity for the H-2L ^d molecule: implications for vaccine design and immunotherapy," <i>Mol. Immunol.</i> 34:273-281, Elsevier Science, Ltd. (1997)

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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>15</u>	Pamer, E.G., et al., "Precise prediction of a dominant class I MHC-restricted epitome of <i>Listeria monocytogenes</i> ," <i>Nature</i> 353:852-855, Macmillan Publishers, Ltd. (1991)
	AS	<u>15</u>	Parham, P., et al., "The Origins of HLA-A,B,C Polymorphism," <i>Immunol. Rev.</i> 143:141-180, Munksgaard (1995)
	AT	<u>15</u>	Parker, K.C., et al., "Peptide Binding to HLA-A2 and HLA-B27 Isolated from <i>Escherichia coli</i> ," <i>J. Biol. Chem.</i> 267:5451-5459, American Society for Biochemistry and Molecular Biology, Inc. (March 1992)

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	AR	<u>16</u>	Parker, K.C., et al., "Sequence motifs important for peptide binding to the human MHC class I molecule, HLA-A2," <i>J. Immunol.</i> 149:3580-3587, American Association of Immunologists (December 1992)
	AS	<u>16</u>	Rammensee, H.-G., et al., "Peptides Naturally Presented by MHC Class I Molecules," <i>Annu. Rev. Immunol.</i> 11:213-244, Annual Reviews, Inc. (January 1993)
	AT	<u>16</u>	Rammensee, H.-G., et al., "MHC ligands and peptide motifs: first listing," <i>Immunogenetics</i> 41:178-228, Springer-Verlag (1995)

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	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>18</u>	Röttschke, O., et al., "Isolation and analysis of naturally processed viral peptides as recognized by cytotoxic T cells," <i>Nature</i> 348:252-254, Macmillan Publishers, Ltd. (1990)
	AS	<u>18</u>	Röttschke, O., et al., "Characterization of Naturally Occurring Minor Histocompatibility Peptides Including H-4 and H-Y," <i>Science</i> 249:283-287, American Association for the Advancement of Science (1990)
	AT	<u>18</u>	Röttschke, O., and Falk, K., "Naturally-occurring peptide antigens derived from the MHC class-I-restricted processing pathway," <i>Immunol. Today</i> 12:447-455, Elsevier Science Publishers, Ltd. (1991)

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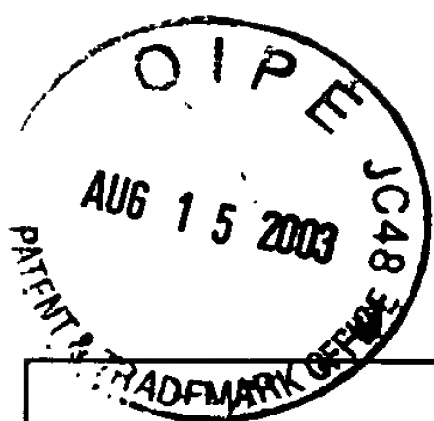
OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>19</u>	Rotzschke, O., et al., "Peptide motifs of closely related HLA class I molecules encompass substantial differences," <i>Eur. J. Immunol.</i> 22:2453-2456, VCH Verlagsgesellschaft mbH (September 1992)
	AS	<u>19</u>	Röttschke, O., and Falk, K., "Origin, structure and motifs of naturally processed MHC class II ligands," <i>Curr. Opin. Immunol.</i> 6:45-51, Current Biology, Ltd. (February 1994)
	AT	<u>19</u>	Rudinger, J., "Characteristics of the amino acids as components of a peptide hormone sequence," in <i>Peptide Hormones</i> , Parsons, J.A., ed., University Park Press, Baltimore, MD, pp.1-7 (1976)

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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>20</u>	Ruppert, J., et al., "Prominent Role of Secondary Anchor Residues in Peptide Binding to HLA-A2.1 Molecules," Cell 74:929-937, Cell Press (September 1993)
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	AT	<u>20</u>	Sette, A., et al., "Prediction of major histocompatibility complex binding regions of protein antigens by sequence pattern analysis," Proc. Natl. Acad. Sci. USA 86:3296-3300, National Academy Press (1989)

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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>21</u>	Sette, A., et al., "Random association between the peptide repertoire of A2.1 class I and several different DR class II molecules," <i>J. Immunol.</i> 147:3893-3900, The American Association of Immunologists (1991)
	AS	<u>21</u>	Sette, A., et al., "The Relationship Between Class I Binding Affinity and Immunogenicity of Potential Cytotoxic T Cell Epitopes," <i>J. Immunol.</i> 153:5586-5592, The American Association of Immunologists (December 1994)
	AT	<u>21</u>	Shastri, N., et al., "Presentation of Endogenous Peptide/MHC Class I Complexes Is Profoundly Influenced by Specific C-Terminal Flanking Residues," <i>J. Immunol.</i> 155: 4339-4346, The American Association of Immunologists (1995)

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	AR	22	Sherman, L.A., et al., "Extracellular Processing of Peptide Antigens That Bind Class I Major Histocompatibility Molecules," <i>J. Exp. Med.</i> 175:1221-1226, The Rockefeller University Press (May 1992)
	AS	22	Shimojo, N., et al., "Specificity of peptide binding by the HLA-A2.1 molecule," <i>J. Immunol.</i> 143:2939-2947, The American Association of Immunologists (1989)
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	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>23</u>	Threlkeld, S.C., et al., "Degenerate and Promiscuous Recognition by CTL of Peptides Presented by the MHC Class I A3-like Superfamily. Implications for Vaccine Development," <i>J. Immunol.</i> 159:1648-1657, The American Association of Immunologists (1997)
	AS	<u>23</u>	Wentworth, P.A., et al., "Differences and similarities in the A2.1-restricted cytotoxic T cell repertoire in humans and human leukocyte antigen-transgenic mice," <i>Eur. J. Immunol.</i> 26:97-101, VCH Verlagsgesellschaft mbH (1996)
	AT	<u>23</u>	Whitton, J.L., et al., "Molecular Analyses of a Five-Amino-Acid Cytotoxic T-Lymphocyte (CTL) Epitope: an Immunodominant Region Which Induces Nonreciprocal CTL Cross-Reactivity," <i>J. Virol.</i> 63:4303-4310, American Society for Microbiology (1989)

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	AR	<u>24</u>	Yewdell, J.W., and Bennink, J.R., "Cell Biology of Antigen Processing and Presentation to Major Histocompatibility Complex Class I Molecule-Restricted T Lymphocytes," <i>Adv. Immunol.</i> 52:1-123, Academic Press (July 1992)
	AS	<u>24</u>	York, I.A., and Rock, K.L., "Antigen processing and presentation by the class I major histocompatibility complex," <i>Annu. Rev. Immunol.</i> 14:369-396, Annual Reviews, Inc. (1996)
	AT	<u>24</u>	Zhang, Q-J., et al., "An HLA-A11-specific motif in nonamer peptides derived from viral and cellular proteins," <i>Proc. Natl. Acad. Sci. USA</i> 90:2217-2221, National Academy Press (March 1993)

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	AR	<u>25</u>	Parker, K.C., et al., "Scheme for Ranking Potential HLA-A2 Binding Peptides Based on Independent Binding of Individual Peptide Side-Chains," <i>J. Immunol.</i> 152:163-175, The American Association of Immunologists (January 1994)
	AS	<u>25</u>	Dialog File 351, Accession No. 7180926, Derwent WPI English language abstract for EP 0 226 513.
	AT	<u>25</u>	Dialog File 351, Accession No. 9263567, Derwent WPI English language abstract for WO 92/21033.

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	AR	<u>26</u>	Dialog File 351, Accession No. 9888606, Derwent WPI English language abstract for WO 94/11738.
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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of:

FIKES *et al.*

Appl. No. 09/458,297

Filed: December 10, 1999

For: **Inducing Cellular Immune
Responses To p53 Using Peptide
and Nucleic Acid Compositions**

Confirmation No. 8696

Art Unit: 1644

Examiner: Schwadron, R.B.

Atty. Docket:
2060.0120000/EKS/HCC/M-M

Third Supplemental Information Disclosure Statement

Commissioner for Patents
PO Box 1450
Alexandria, VA 22313-1450

Sir:

Listed on accompanying Form PTO-1449 are documents that may be considered material to the examination of this application, in compliance with the duty of disclosure requirements of 37 C.F.R. §§ 1.56, 1.97 and 1.98. The numbering on this Third Supplemental Information Disclosure Statement is a continuation of the numbering in Applicants' Second Supplemental Information Disclosure Statement filed herewith in connection with the above-captioned application. A copy of each document is also provided.

The Examiner's attention is directed to the following co-pending U.S. patent applications:

U.S. Application No. 08/205,713, filed March 4, 1994;

U.S. Application No. 08/344,824, filed November 23, 1994;

U.S. Application No. 08/347,610, filed December 1, 1994;

U.S. Application No. 08/591,502, filed May 20, 1996;

U.S. Application No. 09/017,524, filed February 3, 1998;

U.S. Application No. 09/665,510, filed September 19, 2000;

U.S. Application No. 10/149,140, § 371 date November 18, 2002;

which are directed to related technical subject matter. The identification of these U.S. patent applications is not to be construed as a waiver of secrecy as to these applications now or upon issuance of the present application as a patent. The Examiner is respectfully requested to consider the cited applications and the art cited therein during examination.

Where the publication date of a listed document does not provide a month of publication, the year of publication of the listed document is sufficiently earlier than the effective U.S. filing date and any foreign priority date so that the month of publication is not in issue. Applicants have listed publication dates on the attached PTO-1449 based on information presently available to the undersigned. However, the listed publication dates should not be construed as an admission that the information was actually published on the date indicated.

Applicants reserve the right to establish the patentability of the claimed invention over any of the information provided herewith, and/or to prove that this information may not be prior art, and/or to prove that this information may not be enabling for the teachings purportedly offered.

This statement should not be construed as a representation that a search has been made, or that information more material to the examination of the present patent application does not exist. The Examiner is specifically requested not to rely solely on the material submitted herewith.

This Third Supplemental Information Disclosure Statement is being filed before the

mailing date of a first Office Action on the merits. No statement or fee is required.

It is respectfully requested that the Examiner initial and return a copy of the enclosed PTO-1449, and indicate in the official file wrapper of this patent application that the documents have been considered.

The U.S. Patent and Trademark Office is hereby authorized to charge any fee deficiency, or credit any overpayment, to our Deposit Account No. 19-0036.

Respectfully submitted,

STERNE, KESSLER, GOLDSTEIN & FOX P.L.L.C.



Helene C. Carlson
Agent for Applicants
Registration No. 47,473

Date: 8/15/03

1100 New York Avenue, N.W.
Washington, D.C. 20005-3934
(202) 371-2600



FORM PTO-1449

THIRD SUPPLEMENTAL
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	AN						Yes No
	AO2	WO 01/41788 A1	06/14/2001	WIPO			Yes No
	AP						Yes No

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	AS	26	Boon, T., "Tumor Antigens Recognized by Cytolytic T Lymphocytes: Present Perspectives for Specific Immunotherapy," <i>Int. J. Cancer</i> 54:177-180, Wiley-Liss, Inc. (May 1993).
	AT	26	Harlow, E., et al., "Molecular Cloning and In Vitro Expression of a cDNA Clone for Human Cellular Tumor Antigen p53," <i>Mol. Cell. Biol.</i> 5:1601-1610, American Society for Microbiology (1985).

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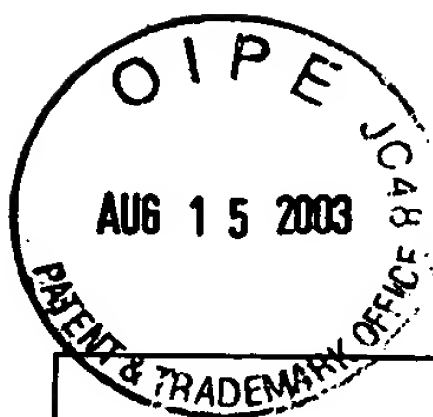
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	AL						Yes No
	AM						Yes No
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	AP						Yes No

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	AR	27	Harris, N., et al., "Molecular Basis for Heterogeneity of the Human p53 Protein," <i>Mol. Cell. Biol.</i> 6:4650-4656, American Society for Microbiology (1986).
	AS	27	Kawashima, I., et al., "The Multi-epitope Approach for Immunotherapy for Cancer: Identification of Several CTL Epitopes from Various Tumor-Associated Antigens Expressed on Solid Epithelial Tumors," <i>Hum. Immunol.</i> 59:1-14, American Society for Histocompatibility and Immunogenetics (1998).
	AT	27	Knuth, A., et al., "Cellular and humoral immune responses against cancer: implications for cancer vaccines," <i>Curr. Opin. Immunol.</i> 3:659-664, Current Biology Ltd. (1991).

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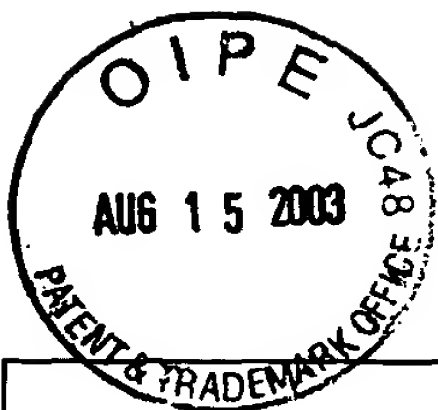
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	AR	<u>28</u>	Lamb, P., and Crawford, L., "Characterization of the Human p53 Gene," <i>Mol. Cell. Biol.</i> 6:1379-1385, American Society for Microbiology (1986).
	AS	<u>28</u>	"Neoplasia," in <i>Robbins Pathologic Basis of Disease</i> , 4 th edition, Cotran, R.S., et al., eds., W.B. Saunders Company, Philadelphia, pp. 296-299 (1989).
	AT	<u>28</u>	Toes, R.E.M., et al., "Enhancement of Tumor Outgrowth Through CTL Tolerization After Peptide Vaccination Is Avoided by Peptide Presentation on Dendritic Cells," <i>J. Immunol.</i> 160:4449-4456, The American Association of Immunologists (1998).

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	AR	<u>29</u>	Zakut-Houri, R., et al., "Human p53 cellular tumor antigen: cDNA sequence and expression in COS cells," <i>EMBO J</i> 4:1251-1255, IRL Press Limited (1985).
	AS	<u>29</u>	Gnjatic, S., et al., "Mapping and ranking of potential cytotoxic T epitopes in the p53 protein: effect of mutations and polymorphism on peptide binding to purified and refolded HLA molecules," <i>Eur. J. Immunol.</i> 25:1638-1642, VCH Verlagsgesellschaft mbH (1995).
	AT	<u>29</u>	Röpke, M., et al., "T Cell-Mediated Cytotoxicity Against p53-Protein Derived Peptides in Bulk and Limiting Dilution Cultures of Healthy Donors," <i>Scand. J. Immunol.</i> 42:98-103, Blackwell Science Limited (1995).

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	AR	<u>30</u>	Vojtesek, B., et al., "Conformational changes in p53 analysed using new antibodies to the core DNA binding domain of the protein," <i>Oncogene</i> 10:389-393, Stockton Press (1995).
	AS	<u>30</u>	Copy of co-pending U.S. application 08/205,713, inventor Grey, filed March 4, 1994 (NOT PUBLISHED).
	AT	<u>30</u>	Copy of co-pending U.S. application 08/344,824, inventors Sette et al., filed November 23, 1994 (NOT PUBLISHED).

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	AR	<u>31</u>	Copy of co-pending U.S. application 08/347,610, inventors Kubo et al., filed December 1, 1994 (NOT PUBLISHED).
	AS	<u>31</u>	Copy of co-pending U.S. application 08/591,502, inventors Chisari et al., filed May 20, 1996 (NOT PUBLISHED).
	AT	<u>31</u>	Copy of co-pending U.S. application 09/017,524, inventors Kubo et al., filed February 3, 1998 (NOT PUBLISHED).

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	AR	32	Copy of co-pending U.S. application 09/665,510, inventors Kubo et al., filed September 19, 2000 (NOT PUBLISHED).
	AS	32	Copy of co-pending U.S. application 10/149,140, inventors Fikes et al., § 371 date November 18, 2002 (PUBLISHED AS WO 01/41788 A1, See Document AO2).
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